- d. Pro at position 217 is replaced by any naturally occurring amino acid other than Pro;
- e. Arg at position 218 is replaced by any naturally occurring amino acid other than Arg;
- f. Ala at position 219 is replaced by any naturally occurring amino acid other than Ala;
- g. Gly at position 220 is replaced by any naturally occurring amino acid other than Gly;
- h. Arg at position 221 is replaced by any naturally occurring amino acid other than Arg;
- i. Ala at position 222 is replaced by any naturally occurring amino acid other than Ala.
- 3. (Amended) A FLINT analog as in Claim [1]  $\underline{2}$  comprising an amino acid substitution in the region defined by amino acids 214 through 222 of SEQ ID NO:1, selected from the group consisting of:
  - a. Gly at position 214 is replaced by a positively charged amino acid that is not Gly;
  - b. Pro at position 215 is replaced by a positively charged amino acid that is not Pro;
  - c. Thr at position 216 is replaced by a positively charged amino acid that is not Thr;
  - d. Pro at position 217 is replaced by a positively charged amino acid that is not Pro;
  - e. Arg at position 218 is replaced by a positively charged amino acid that is not Arg;
  - f. Ala at position 219 is replaced by a positively charged amino acid that is not Ala;
  - g. Gly at position 220 is replaced by a positively charged amino acid that is not Gly;
  - h. Arg at position 221 is replaced by a positively charged amino acid that is not Arg; or

- i. Ala at position 222 is replaced by a positively charged amino acid that is not Ala.
- 4. (Amended) A FLINT analog as in Claim [1]  $\underline{2}$  comprising an amino acid substitution in the region defined by amino acids 214 222 of SEQ ID NO:1, selected from the group consisting of:
  - a. Gly at position 214 is replaced by a negatively charged amino acid that is not Gly;
  - b. Pro at position 215 is replaced by a negatively charged amino acid that is not Pro;
  - c. Thr at position 216 is replaced by a negatively charged amino acid that is not Thr;
  - d. Pro at position 217 is replaced by a negatively charged amino acid that is not Pro;
  - e. Arg at position 218 is replaced by a negatively charged amino acid that is not Arg;
    - f. Ala at position 219 is replaced by a negatively charged amino acid that is not Ala;
    - g. Gly at position 220 is replaced by a negatively charged amino acid that is not Gly;
      - h. Arg at position 221 is replaced by a negatively charged amino acid that is not Arg; or
      - i. Ala at position 222 is replaced by a negatively charged amino acid that is not Ala.
- 5. (Amended) A FLINT analog as in Claim [1]  $\underline{2}$  comprising an amino acid substitution in the region defined by amino acids 214 222 of SEQ ID NO:1, selected from the group consisting of:
  - a. Gly at position 214 is replaced by a polar uncharged amino acid that is not Gly;

- b. Pro at position 215 is replaced by a polar uncharged amino acid that is not Pro;
- c. Thr at position 216 is replaced by a polar uncharged amino acid that is not Thr;
- d. Pro at position 217 is replaced by a polar uncharged amino acid that is not Pro;
- e. Arg at position 218 is replaced by a polar uncharged amino acid that is not Arg;
- f. Ala at position 219 is replaced by a polar uncharged amino acid that is not Ala;
- g. Gly at position 220 is replaced by a polar uncharged amino acid that is not Gly;
- h. Arg at position 221 is replaced by a polar uncharged amino acid that is not Arg; or
- i. Ala at position 222 is replaced by a polar uncharged amino acid that is not Ala.
- 6. (Amended) A FLINT analog as in Claim [1]  $\underline{2}$  comprising an amino acid substitution in the region defined by amino acids 214 222 of SEQ ID NO:1, selected from the group consisting of:
  - a. Gly at position 214 is replaced by a nonpolar amino acid that is not Gly;
  - b. Pro at position 215 is replaced by a nonpolar amino acid that is not Pro;
  - c. Thr at position 216 is replaced by a nonpolar amino acid that is not Thr;
  - d. Pro at position 217 is replaced by a nonpolar amino acid that is not Pro;
  - e. Arg at position 218 is replaced by a nonpolar amino acid that is not Arg;
  - f. Ala at position 219 is replaced by a nonpolar amino acid that is not Ala;

- g. Gly at position 220 is replaced by a nonpolar amino acid that is not Gly;
- h. Arg at position 221 is replaced by a nonpolar amino acid that is not Arg; or
- i. Ala at position 222 is replaced by a nonpolar amino acid that is not Ala.
- 7. (Amended) A FLINT analog [as in Claim 1] resistant to proteolysis at position 218 of SEQ ID NO:1 comprising an amino acid substitution in SEQ ID NO:1, selected from the group consisting of:
  - a.Arg at position 218 is replaced by Gln;
  - b. Arg at position 218 is replaced by Glu;
  - c. Thr at position 216 is replaced by Pro;
  - d. Arg at position 218 is replaced by Ala;
  - e. Arg at position 218 is replaced by Gly;
  - f. Arg at position 218 is replaced by Ser;
  - g. Arg at position 218 is replaced by Val
  - h. Arg at position 218 is replaced by Tyr;
  - i. Arg at position 218 is replaced by Asn; and
  - j. Pro at position 217 is replaced by Tyr.
- 14. (Amended) A method to treat or prevent a disease or condition in a mammal comprising the administration of a therapeutically-effective amount of a protease resistant FLINT analog of Claim [1-13] 2.
- 16. (Amended) A pharmaceutical formulation comprising as an active ingredient a protease resistant FLINT analog of Claim [1-13] 2 associated with one or more pharmaceutically acceptable carriers, excipients, or diluents thereof.

COVERT CTOURS

A2